



FOLEY & LARDNER LLP
ATTORNEYS AT LAW
WASHINGTON HARBOUR
3000 K STREET, N.W., SUITE 500
WASHINGTON, D.C. 20007-5143
TELEPHONE: 202.672.5300
FACSIMILE: 202.672.5399
WWW.FOLEY.COM

FACSIMILE TRANSMISSION

Total # of Pages **4** (including this page)

| TO: | PHONE #: | FAX #: |
|---------------------------|----------------|----------------|
| USPTO Doretha Marcelli | (703) 305-4762 | (703) 308-7333 |

From : Stephen A. Bent
Email Address : sbent@foley.com
Sender's Direct Dial : 202.672.5404
Date : October 21, 2004
Client/Matter No : 023001-0102
User ID No : 0056

MESSAGE:

Attached, per your request, is a copy of the SB08 filed June 21, 2002. Please note that our reference number for this matter changed, from 051023-0115 to **023001-0102!**

Thanks so much!

If there are any problems with this transmission or if you have not received all of the pages, please call 202.672.5340.

| | | |
|-----------|------------|------------------------------------------|
| Operator: | Time Sent: | Return Original To: Camille J. Butler |
|-----------|------------|------------------------------------------|

CONFIDENTIALITY NOTICE: THE INFORMATION CONTAINED IN THIS FACSIMILE MESSAGE IS INTENDED ONLY FOR THE PERSONAL AND CONFIDENTIAL USE OF THE DESIGNATED RECIPIENTS NAMED ABOVE. THIS MESSAGE MAY BE AN ATTORNEY-CLIENT COMMUNICATION, AND AS SUCH IS PRIVILEGED AND CONFIDENTIAL. IF THE READER OF THIS MESSAGE IS NOT THE INTENDED RECIPIENT OR ANY AGENT RESPONSIBLE FOR DELIVERING IT TO THE INTENDED RECIPIENT, YOU ARE HEREBY NOTIFIED THAT YOU HAVE RECEIVED THIS DOCUMENT IN ERROR, AND THAT ANY REVIEW, DISSEMINATION, DISTRIBUTION OR COPYING OF THIS MESSAGE IS STRICTLY PROHIBITED. IF YOU HAVE RECEIVED THIS COMMUNICATION IN ERROR, PLEASE NOTIFY US IMMEDIATELY BY TELEPHONE AND RETURN THE ORIGINAL MESSAGE TO US BY MAIL. THANK YOU.

PTO/SB/08 (08-00)

Approved for use through 10/31/2002. OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

| | | | | | |
|------------------------------------------------------------------------------------------------------------------------------------------------------------------|---|----|---|--------------------------|--------------------|
| Substitute for form 1449B/PTO INFORMATION DISCLOSURE STATEMENT BY APPLICANT Date Submitted: 06/21/2002 (use as many sheets as necessary) | | | | Complete if Known | |
| | | | | Application Number | 10/049,373 |
| | | | | Filing Date | 06/21/2002 |
| | | | | First Named Inventor | Isao ISHIDA et al. |
| | | | | Group Art Unit | Unassigned |
| | | | | Examiner Name | Unassigned |
| | | | | Attorney Docket Number | 051023-0115 |
| Sheet | 1 | of | 3 | | |

| U.S. PATENT DOCUMENTS | | | | | | |
|-----------------------|-----------------------|----------------------|-----------------------------------|-------------------------------------------------|--------------------------------------------------|---------------------------------------------------------------------------|
| Examiner Initials* | Cite No. ¹ | U.S. Patent Document | | Name of Patentee or Applicant of Cited Document | Date of Publication of Cited Document MM-DD-YYYY | Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear |
| | | Number | Kind Code ² (if known) | | | |
| | | | | | | |
| | | | | | | |
| | | | | | | |
| | | | | | | |
| | | | | | | |
| | | | | | | |
| | | | | | | |
| | | | | | | |
| | | | | | | |

| FOREIGN PATENT DOCUMENTS | | | | | | | | |
|--------------------------|-----------------------|-------------------------|---------------------|-----------------------------------|--------------------------------------------------|--------------------------------------------------|---------------------------------------------------------------------------|----------------|
| Examiner Initials* | Cite No. ¹ | Foreign Patent Document | | | Name of Patentee or Applicant of Cited Documents | Date of Publication of Cited Document MM-DD-YYYY | Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear | T ⁶ |
| | | Office ³ | Number ⁴ | Kind Code ⁵ (if known) | | | | |
| | A1 | WO | 98/37757 | A1 | Tomizuka et al. | 3/09/1998 | | abs |
| | | | | | | | | |
| | | | | | | | | |
| | | | | | | | | |
| | | | | | | | | |
| | | | | | | | | |
| | | | | | | | | |
| | | | | | | | | |
| | | | | | | | | |

| OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS | | | |
|---------------------------------------------------|-----------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------|
| Examiner Initials* | Cite No. ¹ | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.) date, page(s), volume-issue number(s), publisher, city and/or country where published. | T ⁶ |
| | A2 | TOMIZUKA et al., EP0773288 A3, July 9, 1997, Abstract. | |
| | A3 | RAMSDEN, R. et al., "Phenobarbital Induction and Tissue-specific Expression of the Rat CYP2B2 Gene in Transgenic Mice", <u>J. Biol. Chem.</u> , 268 (222): 21722-2176 (1993). | |
| | A4 | KAMATAKI, T. et al., "Application of Molecular Technology to Drug Metabolism Studies: Perspectives", <u>Yakubutsu Doutai</u> , 13(3): 280-286 (1997) | yes |
| | A5 | NELSON et al., "P450 superfamily: update on new sequences, gene mapping, accession numbers and nomenclature," <u>Pharmacogenetics</u> , 6: 1-42, (1996). | |

| | | | |
|--------------------|--|-----------------|--|
| Examiner Signature | | Date Considered | |
|--------------------|--|-----------------|--|

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹ Unique citation designation number. ² See attached Kinds of U.S. Patent Documents. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document.

⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

Burden Hour Statement: This form is estimated to take 2.0 hours to complete. Time will vary depending upon the needs of the individual case. Any comments on the amount of time you are required to complete this form should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, Washington, D.C. 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Assistant Commissioner for Patents, Washington, D.C. 20231.

002.642227.1

PTO/SB/08 (08-00)

Approved for use through 10/31/2002. OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

| | | | | | |
|-------------------------------------------------------------------------------------------------------------------------------------|---|----|---|--------------------------|--------------------|
| Substitute for form 1449B/PTO | | | | Complete if Known | |
| INFORMATION DISCLOSURE STATEMENT BY APPLICANT Date Submitted: 06/21/2002 (use as many sheets as necessary) | | | | Application Number | 10/049,373 |
| | | | | Filing Date | 06/21/2002 |
| | | | | First Named Inventor | Isao ISHIDA et al. |
| | | | | Group Art Unit | Unassigned |
| | | | | Examiner Name | Unassigned |
| Sheet | 2 | of | 3 | Attorney Docket Number | 051023-0115 |

| OTHER PRIOR ART – NON PATENT LITERATURE DOCUMENTS | | | | |
|---------------------------------------------------|----------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------|--|
| Examiner Initials* | Cite No. | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.) date, page(s), volume-issue number(s), publisher, city and/or country where published. | T ^d | |
| | A6 | TOMIZUKA, K. <i>et al.</i> , "Functional Expression and Germline Transmission of a Human Chromosome Fragment in Chimaeric Mice", <u>Nature Genetics</u> , 16: 133-143 (1997). | | |
| | A7 | JAKOBOVITS, A. <i>et al.</i> , "Germ-line Transmission and Expression of a Human-Derived Yeast Artificial Chromosome", <u>Nature</u> , 362: 255-258 (1993). | | |
| | A8 | SHIINA, T. <i>et al.</i> , "Nucleotide Sequencing Analysis of the 146-Kilobase Segment Around the <i>IKBL</i> and <i>MICA</i> Genes at the Centromeric End of the HLA Class 1 Region", <u>Genomics</u> , 47: 372-382 (1998). | | |
| | A9 | HUBERT, R. <i>et al.</i> , "BAC and PAC Contigs Covering 3.5 Mb of the Down Syndrome Congenital Heart Disease Region Between D21S55 and MX1 on Chromosome 21", <u>Genomics</u> , 41: 218-226 (1997). | | |
| | A10 | ZHI-WEI, LI <i>et al.</i> , "Generation of Mice with a 200-kb Amyloid Precursor Protein Gene Deletion by Cre Recombinase-Mediated Site-Specific Recombination in Embryonic Stem Cells", <u>Proc. Natl. Acad. Sci. USA</u> , 93: 6158-6161 (1996). | | |
| | A11 | GU, H. <i>et al.</i> , "Deletion of a DNA Polymerase β Gene Segment in T Cells Using Cell Type-Specific Gene Targeting", <u>Science</u> , 265: 103-106 (1994). | | |
| | A12 | LI, Y., <i>et al.</i> , "Establishment of Transgenic Mice Carrying Human Fetus-Specific CYP3A7", <u>Archives of Biochemistry and Biophysics</u> , 239 (2): 235-240 (1996). | | |
| | A13 | KAMATAKI, T. <i>et al.</i> , "Expression of CYP3A7, a Human Fetus-Specific Cytochrome P450, in cultured Cells and in the Hepatocytes of p53-Knockout Mice", <u>Toxicology Letters</u> , 82/83: 879-882 (1995). | | |
| | A14 | KOVALEVA, I. <i>et al.</i> , "Transgenic Yeast Expressing Human Cytochrome P450s Can Serve as a Tool in Studies of the Mechanisms of their Induction by Various Effectors", <u>Biochemical and Biophysical Research Communications</u> , 221(1): 129-132 (1996). | | |
| | A15 | GILLAM, E. <i>et al.</i> , "Expression of Modified Human Cytochrome P450 3A4 in <i>Escherichia coli</i> and Purification and Reconstitution of the Enzyme", <u>Archives of Biochemistry and Biophysics</u> , 305(1): (1993). | | |
| | A16 | ASSEFFA, A. <i>et al.</i> , "Novel Exogenous Heme-Dependent Expression of Mammalian Cytochrome P450 Using Baculovirus", <u>Archives of Biochemistry and Biophysics</u> , 274(2): 481-490 (1989). | | |
| | A17 | SHOU, M. <i>et al.</i> , "The Role of 12 cDNA-Expressed Human, Rodent, and Rabbit Cytochromes P450 in the Metabolism of Benzo[a]pyrene and Benzo[a]pyrene <i>trans</i> , 7-8-Dihydrodiol", <u>Molecular Carcinogenesis</u> , 10: 159-168 (1994). | | |
| | A18 | WOLF, C., <i>et al.</i> , "Use of Transgenic Animals in Understanding Molecular Mechanisms of Toxicity", <u>J. Phar. Pharmacol.</u> , 50: 567-574 (1998). | | |
| | A19 | McKINNON, R. <i>et al.</i> , "Cytochrome P450 Knockout Mice: New Toxicological Models", <u>Clinical and Experimental Pharmacology and Physiology</u> , 25: 783-787 (1998). | | |
| | A20 | PINEAU, T. <i>et al.</i> , "Neonatal Lethality Associated with Respiratory Distress in Mice Lacking Cytochrome P450 1A2", <u>Proc Natl. Acad. Sci. USA</u> , 92: 5134-5138 (1995). | | |

| | |
|--------------------|-----------------|
| Examiner Signature | Date Considered |
|--------------------|-----------------|

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 809. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹ Unique citation designation number. ² See attached Kinds of U.S. Patent Documents. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document.

⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

Burden Hour Statement: This form is estimated to take 2.0 hours to complete. Time will vary depending upon the needs of the individual case. Any comments on the amount of time you are required to complete this form should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, Washington, D.C. 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Assistant Commissioner for Patents, Washington, D.C. 20231.

002.842227.1

PTO/SB/08 (08-00)

Approved for use through 10/31/2002. OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

| | | | | | |
|-------------------------------------------------------------------------------------------------------------------------------------|---|----|---|--------------------------|--------------------|
| Substitute for form 1449B/PTO | | | | Complete if Known | |
| INFORMATION DISCLOSURE STATEMENT BY APPLICANT Date Submitted: 06/21/2002 (use as many sheets as necessary) | | | | Application Number | 10/049,373 |
| | | | | Filing Date | 06/21/2002 |
| | | | | First Named Inventor | Isao ISHIDA et al. |
| | | | | Group Art Unit | Unassigned |
| | | | | Examiner Name | Unassigned |
| Sheet | 3 | of | 3 | Attorney Docket Number | 051023-0115 |

| OTHER PRIOR ART – NON PATENT LITERATURE DOCUMENTS | | | | |
|---------------------------------------------------|-----------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------|--|
| Examiner Initials* | Cite No. ¹ | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.) date, page(s), volume-issue number(s), publisher, city and/or country where published. | T ⁶ | |
| | A21 | LIANG, H. et al., "Cyp1a2(-/-) Null Mutant Mice Develop Normally but Show Deficient Drug Metabolism", <u>Proc. Natl. Acad. Sci. USA</u> , 93: 1671-1676 (1996). | | |
| | A22 | GENTER, M. et al., "Role of CYP2A5 and 2G1 in Acetaminophen Metabolism and Toxicity in the Olfactory Mucosa of the Cyp1a2(-/-) Mouse", <u>Biochemical Pharmacology</u> , 55: 1819-1826 (1998). | | |
| | A23 | LEE, S. et al., "Role of CYP2E1 in the Hepatotoxicity of Acetaminophen", <u>The Journal of Biological Chemistry</u> , 271: 20, 12063-12067 (1996). | | |
| | A24 | FUNAE, Y. et al., "Drug Metabolism by Human Cytochrome P 450", <u>Bioscience and Industry</u> , 55: (2): 81-86 (1997). | partial | |
| | A25 | KAMATAKI, T. et al., "Reports of Anhyo Center Laboratory", <u>Incorporated Foundation</u> , 7: 27-33 (1997). | partial | |
| | A26 | ISHIDA et al., "Introduction to Cell Engineering, Experiment Manipulation", <u>Kodansha Publishing</u> , 39: lines 1-4 (1992). | partial | |
| | A27 | AIZAWA, S. "Gene Targeting", <u>Biomaterial Series 8</u> , Yodosha : 89-105 (1995). | | |
| | A28 | SHIMUZU, M., "III Chromosome and Gene Mapping 7, Method for Introduction of Human Chromosome by Microcell Fusion", <u>Cell Engineering Handbook</u> , Yodosha : 127-132 (1992). | partial | |
| | | | | |

| | | | |
|--------------------|--|-----------------|--|
| Examiner Signature | | Date Considered | |
|--------------------|--|-----------------|--|

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹ Unique citation designation number. ² See attached Kinds of U.S. Patent Documents. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document.

⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

Burden Hour Statement: This form is estimated to take 2.0 hours to complete. Time will vary depending upon the needs of the individual case. Any comments on the amount of time you are required to complete this form should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, Washington, D.C. 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Assistant Commissioner for Patents, Washington, D.C. 20231.

002.842227.1